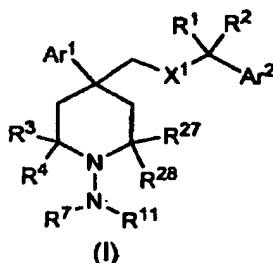


Serial No. 10/612,176
June 29, 2005
Page 2 of 20

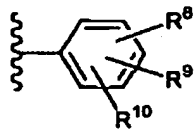
Amendments

Claim 1 (Previously Presented): A compound having the formula (I):



or a pharmaceutically acceptable salt or solvate thereof, wherein

Ar¹ and Ar² are each independently selected from the group consisting of (R¹⁹)_n-heteroaryl- and



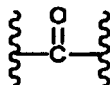
X¹ is selected from the group consisting of -O-, -S-, -SO-, -SO₂-, -NR¹²-, -N(COR¹²)- and -N(SO₂R¹⁵)-;

R¹, R³ and R⁵ are each independently selected from the group consisting of H and C₁-C₈ alkyl;

R², R⁴ and R⁶ are each independently selected from the group consisting of H, -CONR¹³R¹⁴ and -(CH₂)_{n1}-G; wherein G is selected from the group consisting of H, -CF₃, -CHF₂, -CH₂F, -OH, -O-(C₁-C₈)alkyl, -SO₂R¹³, -O-(C₃-C₈ cycloalkyl), -NR¹³R¹⁴, -SO₂NR¹³R¹⁴, -NR¹³SO₂R¹⁵, -NR¹³COR¹², -NR¹²(CONR¹³R¹⁴), -CONR¹³R¹⁴, -COOR¹² and C₃-C₈ cycloalkyl; or

R¹ and R², taken together with the carbon to which they are attached, form a C₃-C₈ cycloalkyl ring; or

R¹ and R², taken together with the carbon to which they are attached, form a

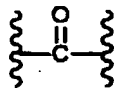


group; or

R³ and R⁴, taken together with the carbon to which they are attached, form a

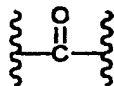
54627-1

Serial No. 10/612,176
June 29, 2005
Page 3 of 20



group; or

R^5 and R^6 , taken together with the carbon to which they are attached, form a



group;

R^7 and R^{11} are each independently selected from the group consisting of H, C_1 - C_8 alkyl, C_3 - C_8 cycloalkyl, $(R^{16})_{n7}$ -aryl-, $(R^{18})_{n7}$ -heteroaryl-, $-COOR^{29}$, $-CONR^{21}R^{22}$, $-CON(R^{21})(CH_2)_n-G^1$, $-S(O)_{n5}(CH_2)_n-G^1$, $-S(O)_{n5}R^{13}$, $-CO(CH_2)_n-G^1$ and $-(CH_2)_{n1}-G^1$; wherein

n is 0-4, and

G^1 is selected from the group consisting of H, $-OH$, $(C_1$ - $C_8)$ alkyl, $-O$ -(C_1 - C_8 alkyl), $-S(O)_{n5}R^{13}$, $-O$ -(C_3 - C_8 cycloalkyl), $-NR^{13}R^{14}$, $-SO_2NR^{13}R^{14}$, $-NR^{13}SO_2R^{15}$, $-NR^{13}COR^{12}$, $-NR^{12}(CONR^{13}R^{14})$, $-OC(=O)R^{12}$, $-CONR^{13}R^{14}$, $-COOR^{12}$, C_3 - C_8 cycloalkyl, $-CF_3$, $(R^{16})_{n7}$ -aryl-O-, $(R^{16})_{n7}$ -aryl-, $(R^{18})_{n7}$ -heteroaryl-, $(R^{19})_{n7}$ -heterocycloalkyl- and alkenyl, and

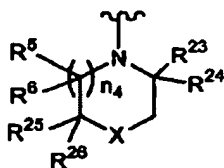
provided that, when n is 0, then G^1 is selected from the group consisting of H, $(C_1$ - $C_8)$ alkyl, alkenyl, $-CONR^{13}R^{14}$, $-COOR^{12}$, C_3 - C_8 cycloalkyl, $-CF_3$, $(R^{16})_{n7}$ -aryl-, $(R^{18})_{n7}$ -heteroaryl-, and $(R^{19})_{n7}$ -heterocycloalkyl-; and

provided that, when n_1 is 1, then G^1 is selected from the group consisting of H, $(C_1$ - $C_8)$ alkyl, alkenyl, $-S(O)_{n5}R^{13}$, $-SO_2NR^{13}R^{14}$, $-CONR^{13}R^{14}$, $-COOR^{12}$, C_3 - C_8 cycloalkyl, $-CF_3$, $(R^{16})_{n7}$ -aryl-, $(R^{18})_{n7}$ -heteroaryl- wherein said heteroaryl ring is bound by a ring carbon to the $-(CH_2)_{n1}-$ group, and $(R^{19})_{n7}$ -heterocycloalkyl- wherein said heterocycloalkyl ring is bound by a ring carbon to the $-(CH_2)_{n1}-$ group; or

R^7 and R^{11} , taken together with the nitrogen to which they are attached, form a 5-7 membered heterocycloalkyl ring of the following formula:

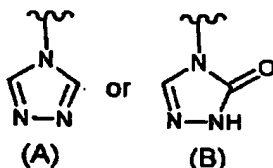
54627-1

Serial No. 10/612,176
 June 29, 2005
 Page 4 of 20



; or

R^7 and R^{11} , taken together with the nitrogen to which they are attached, form a 5-membered ring having the formula (A) or (B):



X is selected from the group consisting of $-NR^{20}$, $-N(CONR^{13}R^{14})$, $-N(CO_2R^{13})$, $-N(SO_2R^{15})$, $-N(COR^{12})$, $-N(SO_2NHR^{13})$, $-O$, $-S$, $-SO$, $-SO_2$, $-CF_2$, $-CH_2$, and $-C(R^{12})F$;

R^8 , R^9 and R^{10} are each independently selected from the group consisting of H, C_1 - C_6 alkyl, C_3 - C_8 cycloalkyl, $-OR^{12}$, halogen, $-CN$, $-NO_2$, $-CF_3$, $-CHF_2$, $-CH_2F$, $-OCF_3$, $-OCHF_2$, $-OCH_2F$, $-COOR^{12}$, $-CONR^{21}R^{22}$, $-NR^{21}COR^{12}$, $-NR^{21}CO_2R^{15}$, $-NR^{21}CONR^{21}R^{22}$, $-NR^{21}SO_2R^{15}$, $-NR^{21}R^{22}$, $-SO_2NR^{21}R^{22}$, $-S(O)_{n5}R^{16}$, $(R^{16})_{n7}$ -aryl- and $(R^{16})_{n7}$ -heteroaryl-;

R^{12} is selected from the group consisting of H, C_1 - C_6 alkyl and C_3 - C_8 cycloalkyl;

R^{13} and R^{14} are each independently selected from the group consisting of H, C_1 - C_6 alkyl, C_2 - C_3 alkyl- $O-CH_3$, C_3 - C_8 cycloalkyl, $(R^{19})_{n7}$ -aryl- $(CH_2)_{n8}$ - and $(R^{19})_{n7}$ -heteroaryl- $(CH_2)_{n8}$ -; or

R^{13} and R^{14} , taken together with the nitrogen to which they are attached, form a 4-7 membered ring containing from 0-3 additional heteroatoms selected from the group consisting of $-O$, $-S$ and $-NR^{12}$;

R^{15} is C_1 - C_6 alkyl, C_3 - C_8 cycloalkyl or $-CF_3$;

R^{16} is 1 to 3 substituents each independently selected from the group consisting of C_1 - C_6 alkyl, C_3 - C_8 cycloalkyl, C_1 - C_6 alkoxy, halogen and $-CF_3$;

R^{19} is 1 to 3 substituents each independently selected from the group consisting of C_1 - C_6 alkyl, C_3 - C_8 cycloalkyl, $-OH$, halogen, $-CN$, $-NO_2$, $-C(O)CF_3$, $-CF_3$, $-CHF_2$, $-CH_2F$, $-OCF_3$, $-OCHF_2$, $-OCH_2F$, $-O-(C_1-C_6 \text{ alkyl})$, $-O-(C_3-C_8 \text{ cycloalkyl})$,

54627-1

Serial No. 10/612,176

June 29, 2005

Page 5 of 20

$-\text{COOR}^{12}$, $-\text{CONR}^{21}\text{R}^{22}$, $-\text{NR}^{21}\text{R}^{22}$, $-\text{NR}^{21}\text{COR}^{12}$, $-\text{NR}^{21}\text{CO}_2\text{R}^{12}$, $-\text{NR}^{21}\text{CONR}^{21}\text{R}^{22}$,
 $-\text{NR}^{21}\text{SO}_2\text{R}^{15}$ and $-\text{S(O)}_{n5}\text{R}^{15}$;

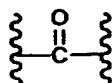
R^{20} is H, $\text{C}_1\text{-C}_6$ alkyl, $\text{C}_3\text{-C}_8$ cycloalkyl, $-(\text{CH}_2)_{n6}\text{-heterocycloalkyl}$,
 $(\text{R}^{19})_{n7}\text{-aryl}(\text{CH}_2)_{n6}\text{-}$ or $(\text{R}^{19})_{n7}\text{-heteroaryl}(\text{CH}_2)_{n6}\text{-}$;

R^{21} and R^{22} are each independently selected from the group consisting of H,
 $\text{C}_1\text{-C}_6$ alkyl, $\text{C}_3\text{-C}_8$ cycloalkyl and benzyl; or

R^{21} and R^{22} , taken together with the nitrogen to which they are attached, form
a 4-7 membered heteroaryl ring containing from 0-3 additional heteroatoms selected
from the group consisting of $-\text{O}-$, $-\text{S}-$ and $-\text{NR}^{12}-$;

R^{23} and R^{24} are each independently selected from the group consisting of H,
 $\text{C}_1\text{-C}_6$ alkyl and $-\text{CONR}^{13}\text{R}^{14}$; or

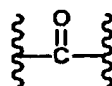
R^{23} and R^{24} , taken together with the carbon atom to which they are attached,
form a



group;

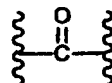
R^{25} , R^{26} , R^{27} and R^{28} are each independently selected from the group
consisting of H and $\text{C}_1\text{-C}_6$ alkyl; or

R^{25} and R^{26} , taken together with the carbon atom to which they are attached,
form a



group; or

R^{27} and R^{28} , taken together with the carbon atom to which they are attached,
form a



group;

R^{29} is selected from the group consisting of $\text{C}_1\text{-C}_6$ alkyl and $\text{C}_3\text{-C}_8$ cycloalkyl;

n_1 is 1-4;

n_4 is 0-2;

n_5 is 0-2;

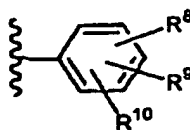
54627-1

Serial No. 10/612,176
June 29, 2005
Page 6 of 20

n_6 is 0-3;
 n_7 is 0-3; and
provided that, when n_4 is 0, and R^{25} and R^{26} are each H, then X is not -O-,
-NR²⁰- or -S-.

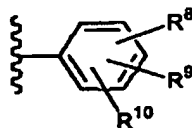
Claim 2 (Original): The compound of Claim 1 wherein X¹ is -O-.

Claim 3 (Original): The compound of Claim 1 wherein Ar¹ and Ar² are each independently



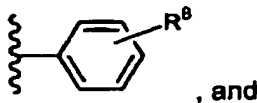
Claim 4 (Original): The compound of Claim 3 wherein R⁸, R⁹ and R¹⁰ are each independently selected from the group consisting of H, -CH₃, halogen and -CF₃.

Claim 5 (Original): The compound of Claim 1 wherein
X¹ is -O-; and
Ar¹ and Ar² are each independently



wherein R⁸, R⁹ and R¹⁰ are each independently selected from the group consisting of H, -CH₃, halogen and -CF₃.

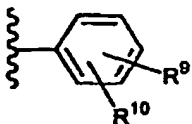
Claim 6 (Original): The compound of Claim 1 wherein
Ar¹ is



Ar² is

54627-1

Serial No. 10/612,176
 June 29, 2005
 Page 7 of 20



wherein,

R^8 is selected from the group consisting of H and F, and

R^9 and R^{10} are each independently selected from the group consisting of H, -CH₃, F, Cl and -CF₃.

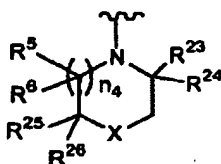
Claim 7 (Original): The compound of Claim 6 wherein

X^1 is -O-; and

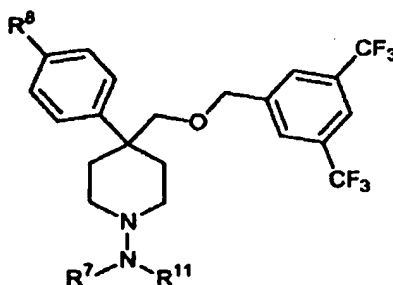
R^3 , R^4 , R^{27} and R^{28} are each H.

Claim 8 (Original): The compound of Claim 6 wherein R^5 and R^6 are H.

Claim 9 (Original): The compound of Claim 7 wherein R^7 and R^{11} , taken together with the nitrogen to which they are attached, form a 5-7 membered ring having the following formula:




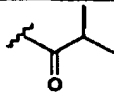
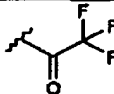
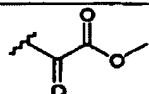
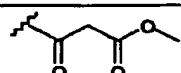
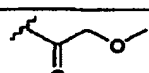
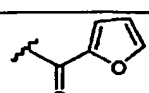
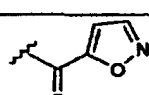
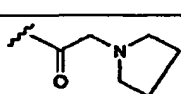
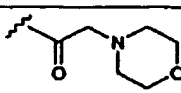
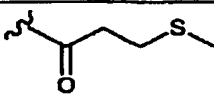
Claim 10 (Original): The compound of Claim 1 having the formula



wherein R^7 , R^8 and R^{11} are selected from the group consisting of:

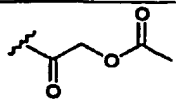
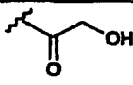
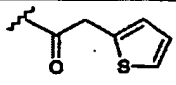
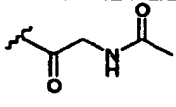
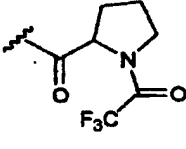
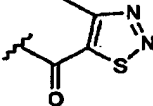
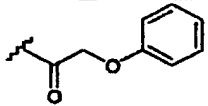


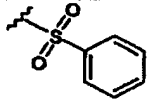
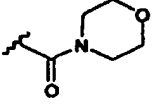
54627-1

Serial No. 10/612,176
June 29, 2005
Page 8 of 20

R ⁸	R ⁷	R ¹¹
H	H	H
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	

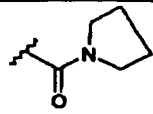
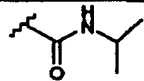
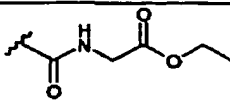
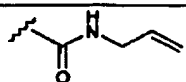
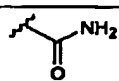
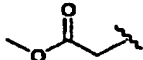
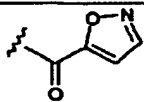
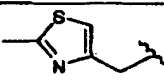
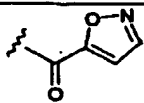
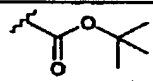
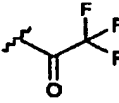
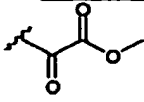
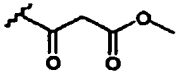
54627-1

Serial No. 10/612,176
June 29, 2005
Page 9 of 20

H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	
H	H	

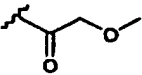
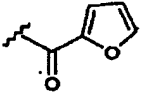
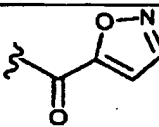
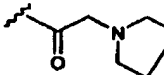
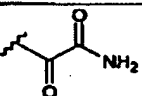
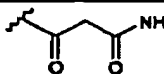
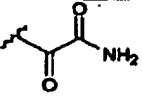
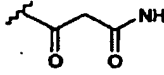
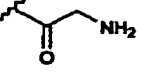
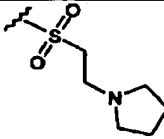
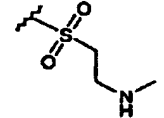
54627-1

Serial No. 10/612,176
June 29, 2005
Page 10 of 20

H	H	
H	H	
H	H	
H	H	
H	H	
H		
H		
F	H	
F	H	H
F	H	
F	H	
F	H	

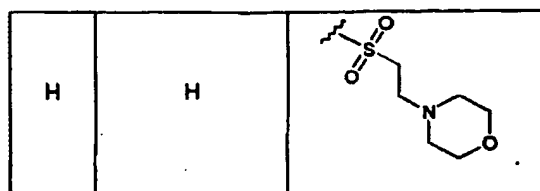
54627-1

Serial No. 10/612,176
June 29, 2005
Page 11 of 20

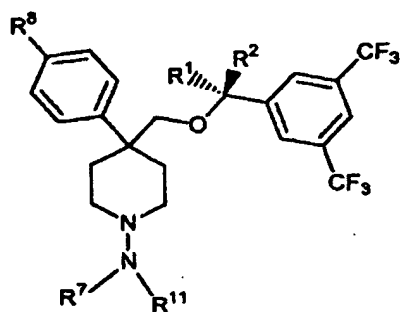
F	H	
F	H	
F	H	
F	H	
H	H	
H	H	
F	H	
F	H	
H	H	
H	H	
H	H	

54627-1

Serial No. 10/612,176
 June 29, 2005
 Page 12 of 20



Claim 11 (Original): The compound of Claim 1 having the formula:

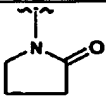
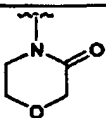
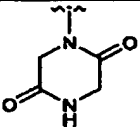
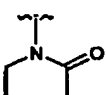
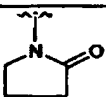
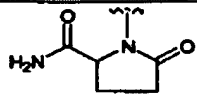
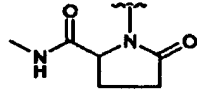
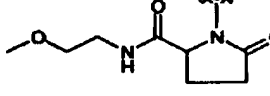
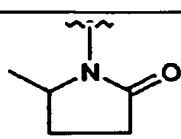
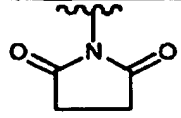


wherein R^1 , R^2 , R^7 , R^8 and R^{11} are selected from the group consisting of:

R^8	R^1	R^2	$-NR^7R^{11}$
H	H	H	
H	H	H	
H	H	H	
H	H	H	
F	H	H	

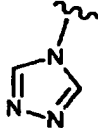
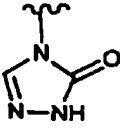
54627-1

Serial No. 10/612,176
June 29, 2005
Page 13 of 20

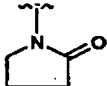
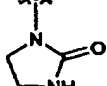
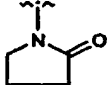
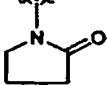
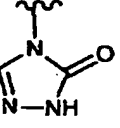
F	H	H	
F	H	H	
H	H	H	
H	H	CH ₃	
H	H	CH ₂ OH	
H	H	H	
H	H	H	
H	H	H	
H	H	H	
H	H	H	

54627-1

Serial No. 10/612,176
June 29, 2005
Page 14 of 20

H	H	H	
H	H	H	

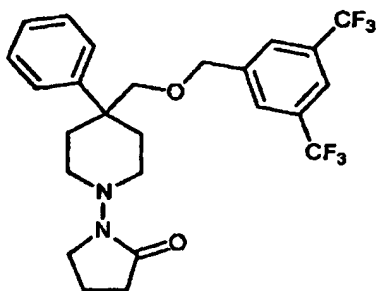
Claim 12 (Original): The compound of Claim 11 or the pharmaceutically acceptable salt or solvate thereof, wherein R^1 , R^2 , R^7 , R^8 and R^{11} are selected from the group consisting of

R^8	R^1	R^2	$-NR^7R^{11}$
H	H	H	
H	H	H	
H	H	CH ₃	
H	H	CH ₂ OH	
H	H	H	

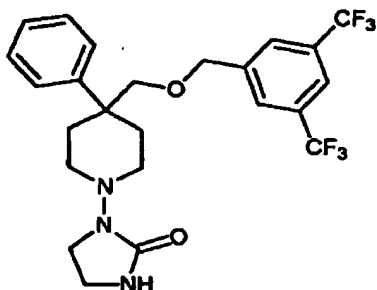
54627-1

Serial No. 10/612,176
June 29, 2005
Page 15 of 20

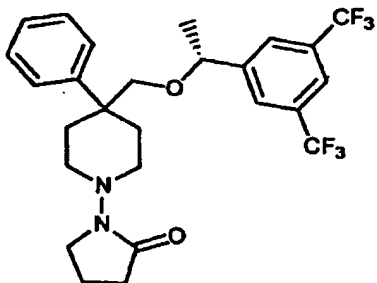
Claim 13 (Original): The compound of Claim 12 having the formula:



Claim 14 (Original): The compound of Claim 12 having the formula:



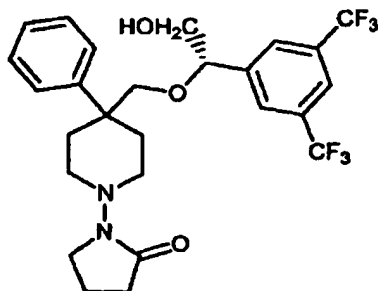
Claim 15 (Original): The compound of Claim 12 having the formula:



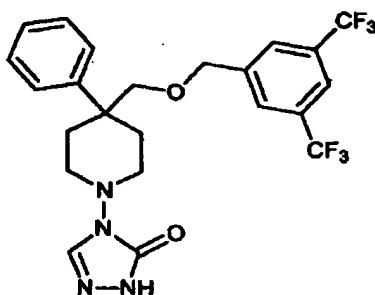
Claim 16 (Original): The compound of Claim 12 having the formula:

54627-1

Serial No. 10/612,176
June 29, 2005
Page 16 of 20



Claim 17 (Original): The compound of Claim 12 having the formula:



Claim 18 (Previously Presented): A pharmaceutical composition comprising a therapeutically effective amount of at least one compound of Claim 1, and a pharmaceutically acceptable carrier.

Claim 19 (Canceled).

Claim 20 (Withdrawn): A method for treating a physiological disorder, symptom or disease in a patient in need of such treatment, comprising administering to said patient an effective amount of at least one compound of Claim 1, wherein said physiological disorder, symptom or disease is selected from the group consisting of: respiratory diseases, inflammatory diseases, skin disorders, ophthalmological disorders, central nervous system conditions, addictions, epilepsy, nociception, psychosis, schizophrenia, Alzheimer's disease, AIDs related dementia, Towne's disease, stress related disorders, obsessive/compulsive disorders, eating disorders, sleep disorders, mania, premenstrual syndrome, gastrointestinal

54627-1

Serial No. 10/612,176
June 29, 2005
Page 17 of 20

disorders, atherosclerosis, fibrosing disorders, obesity, Type II diabetes, pain related disorders, bladder and genitourinary disorders, and nausea.

Claim 21 (Withdrawn): A method for treating a physiological disorder, symptom or disease in a patient in need of such treatment, comprising administering to said patient an effective amount of at least one compound of Claim 1, wherein said physiological disorder, symptom or disease is selected from the group consisting of: a respiratory disease, depression, anxiety, phobia, bipolar disorder, alcohol dependence, psychoactive substance abuse, nociception, psychosis, schizophrenia, stress related disorder, obsessive/compulsive disorder, bulimia, anorexia nervosa, binge eating, sleep disorder, mania, premenstrual syndrome, gastrointestinal disorder, obesity, headache, neuropathic pain, post-operative pain, chronic pain syndrome, bladder disorder, genitourinary disorder, cough, emesis and nausea.

Claim 22 (Withdrawn): A method for treating a physiological disorder, symptom or disease in a patient in need of such treatment, comprising administering to said patient an effective amount of at least one compound of Claim 1, and an effective amount of at least one active ingredient selected from the group consisting of: other NK₁ receptor antagonists, selective serotonin reuptake inhibitors, dopamine receptor agonists, serotonin 5-HT₃ receptor antagonists, serotonin 5-HT_{2c} receptor agonists, nociceptin receptor agonists, glucocorticoids and inhibitors of multidrug resistance protein 5, wherein said physiological disorder, symptom or disease is selected from the group consisting of: a respiratory disease, depression, anxiety, phobia, bipolar disorder, alcohol dependence, psychoactive substance abuse, nociception, psychosis, schizophrenia, stress related disorder, obsessive/compulsive disorder, bulimia, anorexia nervosa, binge eating, sleep disorder, mania, premenstrual syndrome, gastrointestinal disorder, obesity, headache, neuropathic pain, post-operative pain, chronic pain syndrome, bladder disorder, genitourinary disorder, cough, emesis and nausea.

Claim 23 (Currently Amended): A method of treating emesis and/or nausea in a patient in need of such treatment comprising administering to said patient an

54627-1

Serial No. 10/612,176
June 29, 2005
Page 18 of 20

effective amount of at least one compound having the formula (I) of Claim 1 in combination with an effective amount of at least one serotonin 5-HT₃ receptor antagonist and/or at least one glucocorticoid.

Claim 24 (Withdrawn): The method of Claim 23 wherein said serotonin 5-HT₃ receptor antagonist is ondansetron and said glucocorticoid is dexamethasone.

Claim 25 (Withdrawn): The method of Claim 21, wherein the physiological disorder, symptom or disease is emesis, depression, anxiety or cough.

Claim 26 (Withdrawn): The method of Claim 25 wherein the physiological disorder, symptom or disease is depression or anxiety.

Claim 27 (Withdrawn): The method of Claim 26, further comprising administering to the patient an effective amount of at least one anti-depressant agent and/or at least one anti-anxiety agent.

Claim 28 (Withdrawn): The method of Claim 25 wherein depression is being treated and said method further comprises administering to the patient an effective amount of at least one selective serotonin reuptake inhibitor.

Claim 29 (Withdrawn): A method for antagonizing an effect of a Substance P at a neurokinin-1 receptor site or for blocking at least one neurokinin-1 receptor, in a patient in need of such treatment, comprising administering to said patient an effective amount of at least one compound of Claim 1.

Claims 30-35 (Canceled).

54627-1

Art Unit: 1625

Claim 36. A method for treating emesis, depression, anxiety or cough in a patient in need of such treatment comprising administering to said patient an effective amount of at least one compound of claim 1.

Claim 37. A method for treating emesis and/or nausea in a patient in need of such treatment comprising administering to said patient an effective amount of at least one compound of claim 1, in combination with an effective amount of ondansetron and/or with an effective amount of dexamethasone.